

10/060, 789

10/22/07

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NEWS 7 JUL 18 CA/Caplus patent coverage enhanced  
NEWS 8 JUL 26 USPATFULL/USPAT2 enhanced with IPC reclassification  
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NEWS 18 SEP 13 FORIS renamed to SOFIS  
NEWS 19 SEP 13 INPADOCDB enhanced with monthly SDI frequency  
NEWS 20 SEP 17 CA/Caplus enhanced with printed CA page images from 1967-1998  
NEWS 21 SEP 17 Caplus coverage extended to include traditional medicine patents  
NEWS 22 SEP 24 EMBASE, EMBAL, and LEMBASE reloaded with enhancements  
NEWS 23 OCT 02 CA/Caplus enhanced with pre-1907 records from Chemisches Zentralblatt  
NEWS 24 OCT 19 BEILSTEIN updated with new compounds  
  
NEWS EXPRESS 19 SEPTEMBER 2007: CURRENT WINDOWS VERSION IS V8.2, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 19 SEPTEMBER 2007.  
  
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FILE 'HOME' ENTERED AT 11:19:52 ON 22 OCT 2007

=> FIL REGISTRY

COST IN U.S. DOLLARS

SINCE FILE

ENTRY

TOTAL

SESSION

FULL ESTIMATED COST

0.21

0.21

FILE 'REGISTRY' ENTERED AT 11:20:04 ON 22 OCT 2007

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STRUCTURE FILE UPDATES: 19 OCT 2007 HIGHEST RN 951118-42-6

DICTIONARY FILE UPDATES: 19 OCT 2007 HIGHEST RN 951118-42-6

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=> E "XX5"/CN 25

E1	1	XX-88-5/CN
E2	1	XX-88-5 SULFATE/CN
E3	0 -->	XX5/CN
E4	1	XXCC 3/CN
E5	1	XXJ/CN
E6	1	XXL 15/CN
E7	1	XXL 3/CN
E8	1	XXX 1 OIL/CN
E9	1	XXXB/CN
E10	1	XY 1/CN
E11	1	XY 12/CN
E12	1	XY 13/CN
E13	1	XY 13-XY 30 MIXT./CN
E14	1	XY 176/CN
E15	1	XY 1933/CN
E16	1	XY 2/CN
E17	1	XY 2289/CN
E18	1	XY 2289, POLYMER WITH XN 2288/CN
E19	1	XY 27/CN
E20	1	XY 30/CN
E21	1	XY 39 HIGH GLOSS/CN
E22	1	XY 4000/CN
E23	1	XY 4000 ACRYLATE METHACRYLATE/CN
E24	1	XY 4000, 2-METHYL-2-PROPENOATE 2-PROPENOATE/CN
E25	1	XY 4000, POLYMER WITH MEH 7851/CN

=> E "RO-1724"/CN 25

E1	1	RO-1-5155/CN
E2	1	RO-1-6272/CN
E3	0 -->	RO-1724/CN

E4	1	RO-4-1398 HYDROCHLORIDE/CN
E5	1	RO-4-6861 HYDROCHLORIDE/CN
E6	1	RO-ADP/CN
E7	1	RO-ATP/CN
E8	1	RO-C 0C15/CN
E9	1	RO-CILLIN/CN
E10	1	RO-CON/CN
E11	1	RO-CYCLINE/CN
E12	1	RO-NEET/CN
E13	1	RO-NEET 6E/CN
E14	1	RO-PAPAV/CN
E15	1	RO-PEL/CN
E16	1	RO-SA 605/CN
E17	1	RO-W 6602/CN
E18	1	RO/SSA RIBONUCLEOPROTEIN (HUMAN GENE RORET)/CN
E19	1	RO363 OXALATE/CN
E20	1	RO5-1162/CN
E21	1	ROACCUTAN/CN
E22	1	ROACCUTANE/CN
E23	1	ROACH AWAY/CN
E24	1	ROACH KILLER A/CN
E25	1	ROACH PRUFE/CN

=> E "RO1724"/CN 25

E1	1	RO-W 6602/CN
E2	1	RO/SSA RIBONUCLEOPROTEIN (HUMAN GENE RORET)/CN
E3	0 -->	RO1724/CN
E4	1	RO363 OXALATE/CN
E5	1	RO5-1162/CN
E6	1	ROACCUTAN/CN
E7	1	ROACCUTANE/CN
E8	1	ROACH AWAY/CN
E9	1	ROACH KILLER A/CN
E10	1	ROACH PRUFE/CN
E11	1	ROAD CERAM/CN
E12	1	ROADBLOCK/LC7 (CHLORACIDOBACTERIUM THERMOPHILUM CLONE BAC M60-018 J19)/CN
E13	1	ROADBLOCK/LC7 (NITROSOCOCCUS OCEANI STRAIN ATCC 19707)/CN
E14	4	ROADBLOCK/LC7 FAMILY PROTEIN (SALINISPORA TROPICA STRAIN CNB-440)/CN
E15	1	ROADBOND/CN
E16	1	ROADGLAS/CN
E17	1	ROADMENT/CN
E18	1	ROADP/CN
E19	1	ROADPATCH/CN
E20	1	ROALDITE/CN
E21	1	ROAPAS D COLOR D 17/CN
E22	1	ROAPAS D COLOR D 6/CN
E23	1	ROAT 01/CN
E24	1	ROAT 02/CN
E25	1	ROAT 03/CN

=> E "RO-20-1724"/CN 25

E1	1	RO-1-5155/CN
E2	1	RO-1-6272/CN
E3	0 -->	RO-20-1724/CN
E4	1	RO-4-1398 HYDROCHLORIDE/CN
E5	1	RO-4-6861 HYDROCHLORIDE/CN
E6	1	RO-ADP/CN
E7	1	RO-ATP/CN
E8	1	RO-C 0C15/CN
E9	1	RO-CILLIN/CN
E10	1	RO-CON/CN
E11	1	RO-CYCLINE/CN
E12	1	RO-NEET/CN

E13	1	RO-NEET 6E/CN
E14	1	RO-PAPAV/CN
E15	1	RO-PEL/CN
E16	1	RO-SA 605/CN
E17	1	RO-W 6602/CN
E18	1	RO/SSA RIBONUCLEOPROTEIN (HUMAN GENE RORET)/CN
E19	1	RO363 OXALATE/CN
E20	1	RO5-1162/CN
E21	1	ROACCUTAN/CN
E22	1	ROACCUTANE/CN
E23	1	ROACH AWAY/CN
E24	1	ROACH KILLER A/CN
E25	1	ROACH PRUFE/CN

=> E "4-(3-BUTOXY-4-METHOXYBENZYL)-2-IMIDAZOLIDINONE"/CN 25

E1	1	4-(3-BUTOXY-2-HYDROXYPROPYL)-1-OXA-4-AZASPIRO(4.5)DECANE/CN
E2	1	4-(3-BUTOXY-4-METHOXY BENZYL)-2-IMIDAZOLIDINONE/CN
E3	0 -->	4-(3-BUTOXY-4-METHOXYBENZYL)-2-IMIDAZOLIDINONE/CN
E4	1	4-(3-BUTOXY-4-METHOXYBENZYL)-2-IMIDAZOLIDINONE/CN
E5	1	4-(3-BUTOXYPHENYL)-2-CHLOROPYRIMIDINE/CN
E6	1	4-(3-BUTYL-1H-INDOL-2-YL)-6-(PYRIDIN-4-YL)-2H-PYRIDAZIN-3-ONE/CN
E7	1	4-(3-BUTYL-3-(5,5,8,8-TETRAMETHYL-5,6,7,8-TETRAHYDRONAPHTHALEN-2-YL)UREIDO)BENZOIC ACID/CN
E8	1	4-(3-BUTYL-7-ETHYL-2-METHYLPYRROLO(1,2-B)PYRIDAZIN-4-YL)BENZONITRILE/CN
E9	1	4-(3-BUTYLAMINO-4-CHLORO-2,5-DIOXO-3-CYCLOPENTENYLIDENE)-1-BUTYL-2,6-DIMETHYL-1,4-DIHYDROPYRIDINE/CN
E10	1	4-(3-BUTYLAMINO-4-CHLORO-2,5-DIOXO-3-CYCLOPENTENYLIDENE)-2,6-DIPHENYL-1-THIOPYRAN/CN
E11	1	4-(3-BUTYLUREIDO)BENZOIC ACID/CN
E12	1	4-(3-BUTYNYL)-2-PHENYLTHIAZOLE/CN
E13	1	4-(3-BUTYNYL)-3-METHYLANISOLE/CN
E14	1	4-(3-BUTYNYLOXY)-1,2,5-OXADIAZOL-3-AMINE/CN
E15	1	4-(3-BUTYNYLOXY)-6-(2-PROPYNLOXY)PYRIMIDINE/CN
E16	1	4-(3-CARBAMOYL-3,3-DIPHENYLPROPYL)-4-ETHYLMORPHOLINIUM ETHYL SULFATE/CN
E17	1	4-(3-CARBAMOYL-4-(PYRIDIN-3-YL)-1H-PYRROL-1-YL)BUTYLAMINE/CN
E18	1	4-(3-CARBAMOYLPHENOXY)-1-NITROBENZENE/CN
E19	1	4-(3-CARBAMOYLPHENOXY)ANILINE/CN
E20	1	4-(3-CARBAMOYLPHENYL)-7-(1-(TERT-BUTOXYCARBONYL)-5-(PIPERIDINOMETHYL)INDOL-2-YL)ISOINDOLINONE/CN
E21	1	4-(3-CARBAMOYLPHENYL)-7-(1H-5-(PIPERIDINOMETHYL)INDOL-2-YL)ISOINDOLINONE/CN
E22	1	4-(3-CARBAMOYLPIPERAZIN-1-YLMETHYL)-3-TRIFLUOROMETHYLBENZOIC ACID ETHYL ESTER/CN
E23	1	4-(3-CARBAMOYLPIPERIDINO)PIPERIDINE/CN
E24	1	4-(3-CARBAMOYLPROPYL)-2,6-DI-TERT-BUTYLPHENOL/CN
E25	1	4-(3-CARBAZOL-9-YL-2-HYDROXYPROPYL)PIPERAZINE-1-CARBOXYLIC TERT-BUTYL ESTER/CN

=> S E2

L1 1 "4-(3-BUTOXY-4-METHOXY BENZYL)-2-IMIDAZOLIDINONE"/CN

=> DIS L1 1 SQIDE

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DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N:Y

L1 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2007 ACS on STN

RN 29925-17-5 REGISTRY

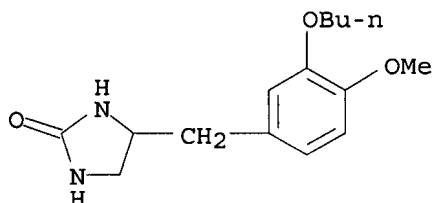
CN 2-Imidazolidinone, 4-[(3-butoxy-4-methoxyphenyl)methyl]- (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 2-Imidazolidinone, 4-(3-butoxy-4-methoxybenzyl)- (8CI)

OTHER NAMES:

CN 4-(3-Butoxy-4-methoxy benzyl)-2-imidazolidinone  
 CN 4-(3-Butoxy-4-methoxybenzyl)-2-imidazolidone  
 CN DL-4-(3-Butoxy-4-methoxybenzyl)-2-imidazolidinone  
 CN R 020-1724  
 CN Ro 20-1724  
 CN Ro 20-174  
 CN Roche 20-1724  
 DR 34185-37-0, 391936-33-7  
 MF C15 H22 N2 O3  
 LC STN Files: AGRICOLA, BEILSTEIN\*, BIOSIS, BIOTECHNO, CA, CAPLUS,  
 CASREACT, CHEMCATS, CIN, CSCHEM, DDFU, DRUGU, EMBASE, IFICDB, IFIPAT,  
 IFIUIDB, IPA, MEDLINE, PHAR, RTECS\*, TOXCENTER, USPAT2, USPATFULL  
 (\*File contains numerically searchable property data)  
 DT.CA CAplus document type: Conference; Journal; Patent  
 RL.P Roles from patents: ANST (Analytical study); BIOL (Biological study);  
 PREP (Preparation); PRP (Properties); RACT (Reactant or reagent); USES  
 (Uses)  
 RLD.P Roles for non-specific derivatives from patents: BIOL (Biological  
 study); USES (Uses)  
 RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological  
 study); PROC (Process); PRP (Properties); RACT (Reactant or reagent);  
 USES (Uses)  
 RLD.NP Roles for non-specific derivatives from non-patents: BIOL (Biological  
 study); PREP (Preparation)



\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

399 REFERENCES IN FILE CA (1907 TO DATE)  
 3 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA  
 399 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> file medline caplus wpids uspatfull  
 COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
8.25	8.46

FULL ESTIMATED COST

FILE 'MEDLINE' ENTERED AT 11:22:05 ON 22 OCT 2007

FILE 'CAPLUS' ENTERED AT 11:22:05 ON 22 OCT 2007

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FILE 'USPATFULL' ENTERED AT 11:22:05 ON 22 OCT 2007

CA INDEXING COPYRIGHT (C) 2007 AMERICAN CHEMICAL SOCIETY (ACS)

=> s 11

L2 875 L1

=> s l2 and (c11 or "chronic lymphocytic leukemia")  
L3 1 L2 AND (CLL OR "CHRONIC LYMPHOCYTIC LEUKEMIA")

=> d l3 ibib, abs, hitstr

L3 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:28040 CAPLUS

DOCUMENT NUMBER: 142:169353

TITLE: Type 4 cAMP phosphodiesterase (PDE4) inhibitors  
augment glucocorticoid-mediated apoptosis in B cell  
chronic lymphocytic leukemia  
(B-CLL) in the absence of exogenous adenylyl  
cyclase stimulation

AUTHOR(S): Tiwari, Sanjay; Dong, Hongli; Kim, Eun Jung;  
Weintraub, Lewis; Epstein, Paul M.; Lerner, Adam

CORPORATE SOURCE: Evans Department of Medicine, Section of Hematology  
and Oncology, Boston Medical Center, Boston, MA,  
02118, USA

SOURCE: Biochemical Pharmacology (2005), 69(3), 473-483  
CODEN: BCPA6; ISSN: 0006-2952

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB CAMP-mediated signaling potentiates glucocorticoid-mediated apoptosis in lymphoid cells, but an effective means by which to take advantage of this observation in the treatment of lymphoid malignancies has not been identified. The primary objective of the current study was to determine whether PDE4 inhibitors, a class of compds. in late clin. development that raise intracellular cAMP levels by inhibiting type 4 cyclic nucleotide phosphodiesterases (PDE4), increase the efficacy of glucocorticoid-mediated apoptosis in leukemic cells from patients with B cell chronic lymphocytic leukemia (B-CLL). Rolipram, a prototypic PDE4 inhibitor, synergized with glucocorticoids in inducing B-CLL but not T cell apoptosis. Rolipram also augmented glucocorticoid receptor element (GRE) transactivation in B-CLL cells. In contrast, inhibition of protein kinase A (PKA) with the cAMP antagonist Rp-8Br-cAMPS reversed both glucocorticoid-induced apoptosis and GRE transactivation. CCRF-CEM cells, a well-studied model of glucocorticoid and cAMP-induced apoptosis, differed from B-CLL cells in that stimulation of adenylyl cyclase with the diterpene forskolin was required to increase both glucocorticoid-mediated apoptosis and GRE activation, while PDE4 inhibition had no effect. Consistent with these results, inhibition of PDE4 induced cAMP elevation in B-CLL but not CCRF-CEM cells, while forskolin augmented cAMP levels in CCRF-CEM but not B-CLL cells. While rolipram treatment up-regulated PDE4B in B-CLL, forskolin treatment up-regulated PDE4D in CCRF-CEM cells. These studies suggest that PKA is required for and enhances glucocorticoid-induced apoptosis in B-CLL by modulating glucocorticoid receptor signal transduction. Clin. trials that examine whether PDE4 inhibitors enhance the efficacy of glucocorticoid-containing chemotherapy regimens in B-CLL are indicated.

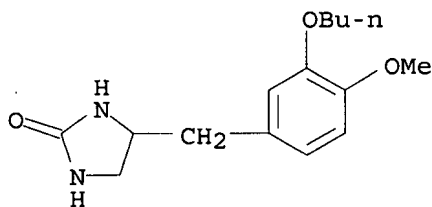
IT 29925-17-5, RO20-1724

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL  
(Biological study); USES (Uses)

(PDE4 inhibitors augment glucocorticoid-mediated apoptosis in B-  
CLL in absence of adenylyl cyclase stimulation)

RN 29925-17-5 CAPLUS

CN 2-Imidazolidinone, 4-[(3-butoxy-4-methoxyphenyl)methyl]- (CA INDEX NAME)



REFERENCE COUNT: 53 THERE ARE 53 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> s l1 and (cancer or tumor)  
L4 140 L1 AND (CANCER OR TUMOR)

=> s l4 and "leukemia"  
L5 30 L4 AND "LEUKEMIA"

=> s l5 and py<2000  
1 FILES SEARCHED...  
L6 7 L5 AND PY<2000

=> d l6 1-7 ibib, abs, hitstr

L6 ANSWER 1 OF 7 MEDLINE on STN  
ACCESSION NUMBER: 97318782 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 9175719  
TITLE: Dissociation between phosphodiesterase inhibition and antiproliferative effects of phosphodiesterase inhibitors on the Dami cell line.  
AUTHOR: Zurbonsen K; Michel A; Vittet D; Bonnet P A; Chevillard C  
CORPORATE SOURCE: INSERM U.300, Montpellier, France.  
SOURCE: Biochemical pharmacology, (1997 Apr 25) Vol. 53, No. 8, pp. 1141-7.  
Journal code: 0101032. ISSN: 0006-2952.  
PUB. COUNTRY: ENGLAND: United Kingdom  
DOCUMENT TYPE: (COMPARATIVE STUDY)  
Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199706  
ENTRY DATE: Entered STN: 30 Jun 1997  
Last Updated on STN: 3 Mar 2000  
Entered Medline: 17 Jun 1997

AB Phosphodiesterase (PDE) inhibitors were shown to inhibit proliferation of various cell types. The present investigation was designed to study the activity of selective PDE inhibitors (8-MeOMIX, milrinone, trequinsin, rolipram, RO-201724, zaprinast, and MY-5445) on the proliferation of the Dami cell line in relation to their effects on cAMP levels and PDE isoenzymes isolated from Dami cells. All compounds, except 8-MeOMIX, elicited antiproliferative effects. Trequinsin, RO-201724, and MY-5445 (100 microM) were found to inhibit cell growth up to 60%, 83%, and 85%, respectively; milrinone, rolipram and zaprinast elicited only weak effects (19-21% at 100 microM). Their growth-inhibitory effects could not be related to their effects on cAMP levels. In addition, although PDE type III and IV inhibitors potentiated cAMP formation due to adenylycyclase activation, no potentiation could be observed when considering their antiproliferative effect. Separation and characterization of PDE of Dami cells revealed the existence of types III, IV, and V isoenzymes. The PDE inhibition found for the PDE inhibitors could not explain their antiproliferative effects. The lack of correlation with cAMP concentrations or PDE inhibition and the high concentrations needed to elicit antiproliferative effects suggest the implication of other

parameters, such as cytotoxicity or lipophilicity, or other targets in addition to PDE for the PDE inhibitors tested. Lipophilicity did not seem to be of importance in antiproliferative effects. In contrast, cytotoxic effects, in particular those of trequinsin and MY-5445, could partially explain their negative action on cell growth.

L6 ANSWER 2 OF 7 MEDLINE on STN  
ACCESSION NUMBER: 97008163 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 8855339  
TITLE: Inhibition of calmodulin-dependent phosphodiesterase induces apoptosis in human leukemic cells.  
AUTHOR: Jiang X; Li J; Paskind M; Epstein P M  
CORPORATE SOURCE: Department of Pharmacology, University of Connecticut Health Center, Farmington 06030, USA.  
SOURCE: Proceedings of the National Academy of Sciences of the United States of America, (1996 Oct 1) Vol. 93, No. 20, pp. 11236-41.  
Journal code: 7505876. ISSN: 0027-8424.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
(RESEARCH SUPPORT, NON-U.S. GOV'T)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
OTHER SOURCE: GENBANK-U56976  
ENTRY MONTH: 199611  
ENTRY DATE: Entered STN: 19 Dec 1996  
Last Updated on STN: 3 Mar 2000  
Entered Medline: 25 Nov 1996

AB Cytosolic extracts from a human lymphoblastoid B-cell line, RPMI-8392, established from a patient with acute lymphocytic leukemia, contain two major forms of cyclic nucleotide phosphodiesterase (PDE): Ca<sup>2+</sup>-calmodulin dependent PDE (PDE1) and cAMP-specific PDE (PDE4). In contrast, normal quiescent human peripheral blood lymphocytes (HPBL) are devoid of PDE1 activity [Epstein, P. M., Moraski, S., Jr., and Hachisu, R. (1987) Biochem. J. 243, 533-539]. Using reverse transcription-polymerase chain reaction (RT-PCR), we show that the mRNA encoding the 63-kDa form of PDE1 (PDE1B1) is expressed in RPMI-8392 cells, but not in normal, resting HPBL. This mRNA is, however, induced in HPBL following mitogenic stimulation by phytohemagglutinin (PHA). Also using RT-PCR, the full open reading frame for human PDE1B1 cDNA was cloned from RPMI-8392 cells and it encodes a protein of 536 amino acids with 96% identity to bovine, rat, and mouse species. RT-PCR also identifies the presence of PDE1B1 in other human lymphoblastoid and leukemic cell lines of B- (RPMI-1788, Daudi) and T- (MOLT-4, NA, Jurkat) cell origin. Inhibition of PDE1 or PDE4 activity by selective inhibitors induced RPMI-8392 cells, as well as the other cell lines, to undergo apoptosis. Culture of RPMI-8392 cells with an 18-bp phosphorothioate antisense oligodeoxynucleotide, targeted against the translation initiation region of the RPMI-8392 mRNA, led to a specific reduction in the amount of PDE1B1 mRNA after 1 day, and its disappearance after 2 days, and induced apoptosis in these cells in a sequence specific manner. This suggests that PDEs, particularly PDE1B1, because its expression is selective, may be useful targets for inducing the death of leukemic cells.

L6 ANSWER 3 OF 7 MEDLINE on STN  
ACCESSION NUMBER: 95061906 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 7971738  
TITLE: Effects of cAMP and cGMP elevating agents on HL-60 cell differentiation.  
AUTHOR: Bang B E; Ericson C; Aarbakke J  
CORPORATE SOURCE: Department of Pharmacology, University of Tromso, Norway.  
SOURCE: Pharmacology & toxicology, (1994 Aug) Vol. 75, No. 2, pp. 108-12.  
Journal code: 8702180. ISSN: 0901-9928.  
PUB. COUNTRY: Denmark



DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
(RESEARCH SUPPORT, NON-U.S. GOV'T)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199412  
ENTRY DATE: Entered STN: 10 Jan 1995  
Last Updated on STN: 10 Jan 1995  
Entered Medline: 6 Dec 1994

AB Previous studies have demonstrated low percentage of HL-60 cell differentiation with theophylline. The present study demonstrate that millimolar concentrations of the non-selective phosphodiesterase inhibitors theophylline, caffeine and isobutyl-methylxanthine all inhibit growth, induce substantial differentiation and elevation of both cAMP and cGMP in HL-60 cells. Selective inhibition of cAMP hydrolysis by Ro20-1724 was without effect. The guanylate cyclase stimulator sodium nitroprusside, which increased cGMP only poorly and also increased cAMP, produced growth inhibition but no differentiation. We put forward the hypothesis that elevation of both cAMP and cGMP above a critical level is necessary for significant cyclic nucleotide induced HL-60 cell differentiation.

L6 ANSWER 4 OF 7 MEDLINE on STN  
ACCESSION NUMBER: 90137009 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 2559336  
TITLE: Histamine inhibits activation of human neutrophils and HL-60 leukemic cells via H2-receptors.  
AUTHOR: Burde R; Seifert R; Buschauer A; Schultz G  
CORPORATE SOURCE: Institut fur Pharmakologie, Freie Universitat Berlin.  
SOURCE: Naunyn-Schmiedeberg's archives of pharmacology, (1989 Dec) Vol. 340, No. 6, pp. 671-8.  
Journal code: 0326264. ISSN: 0028-1298.  
PUB. COUNTRY: GERMANY, WEST: Germany, Federal Republic of  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
(RESEARCH SUPPORT, NON-U.S. GOV'T)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199003  
ENTRY DATE: Entered STN: 28 Mar 1990  
Last Updated on STN: 3 Feb 1997  
Entered Medline: 12 Mar 1990

AB The effects of prostaglandin E1 (PGE1) and histamine on activation of superoxide (O2-) formation, exocytosis of beta-glucuronidase and aggregation in human neutrophils and HL-60 leukemic cells were studied. PGE1, histamine and impromidine, a potent H2-agonist, inhibited O2- formation in neutrophils induced by the chemotactic peptide, N-formyl-L-methionyl-L-leucyl-L-phenylalanine (fMet-Leu-Phe) with IC50 values of 0.5 microM, 8 microM and 2 microM, respectively. The full H1-agonist and weak partial H2-agonist, betahistine, was much less potent and effective than histamine. Dibutyryl cyclic AMP and forskolin mimicked the effects of histamine and PGE1 on O2- formation. The H2-antagonist, famotidine, competitively reversed histamine-induced inhibition of O2- formation with a pA2 value of 7.5. Histamine inhibited O2- formation when added prior to or after fMet-Leu-Phe. fMet-Leu-Phe-induced aggregation and release of beta-glucuronidase in neutrophils were less sensitive to inhibition by PGE1, histamine, dibutyryl cyclic AMP and forskolin than O2- formation. The inhibitor of cyclic AMP-specific phosphodiesterase, rac-4-(3-butoxy-4-methoxybenzyl)-2-imidazolidinone (Ro 20-1724), additively enhanced the inhibitory effects of histamine and PGE1 on the above cell functions. In HL-60 cells differentiated by dimethyl sulfoxide or dibutyryl cyclic AMP, histamine, impromidine and PGE1 but not betahistine inhibited fMet-Leu-Phe-induced O2- formation as well. Our data suggest that histamine inhibits activation of neutrophils and HL-60 cells via H2-receptors through activation of adenylyl cyclase and increased formation of cyclic AMP. (ABSTRACT TRUNCATED AT 250 WORDS)

L6 ANSWER 5 OF 7 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2003:1227 CAPLUS  
DOCUMENT NUMBER: 138:66667  
TITLE: Methods for identifying compounds for inhibiting of  
neoplastic lesions, and pharmaceutical compositions  
containing such compounds  
INVENTOR(S): Pamukcu, Rifat; Piazza, Gary A.  
PATENT ASSIGNEE(S): Cell Pathways, Inc., USA  
SOURCE: U.S., 53 pp., Cont.-in-part of U. S. Ser. No. 46,739.  
CODEN: USXXAM  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 2  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6500610	B1	20021231	US 1999-414625	19991008
US 5858694	A	19990112	US 1997-866027	19970530 <--
CA 2238283	A1	19981130	CA 1998-2238283	19980520 <--
CA 2238283	C	20020820		
TW 591111	B	20040611	TW 1998-87108072	19980525
CZ 295868	B6	20051116	CZ 1998-1651	19980528
NO 9802477	A	19981201	NO 1998-2477	19980529 <--
NO 321717	B1	20060626		
AU 9869794	A	19981210	AU 1998-69794	19980529 <--
AU 709666	B2	19990902		
JP 11094823	A	19990409	JP 1998-150033	19980529 <--
JP 3053381	B2	20000619		
ZA 9804646	A	19991129	ZA 1998-4646	19980529 <--
JP 2000198746	A	20000718	JP 2000-44184	19980529
AT 198771	T	20010215	AT 1998-304247	19980529
ES 2132055	T3	20010501	ES 1998-304247	19980529
IL 124699	A	20030212	IL 1998-124699	19980529
CN 1224761	A	19990804	CN 1998-102044	19980601 <--
CN 1122110	B	20030924		
HK 1012196	A1	20010412	HK 1998-113546	19981216
US 6156528	A	20001205	US 1998-216070	19981219
JP 2000028601	A	20000128	JP 1999-189615	19990702
JP 3234818	B2	20011204		
US 2003004093	A1	20030102	US 2002-40776	20020107
US 2003064421	A1	20030403	US 2002-253849	20020924
US 2003190686	A1	20031009	US 2002-252983	20020924

PRIORITY APPLN. INFO.:

US 1997-866027	A2 19970530
US 1998-46739	A2 19980324
JP 1998-150033	A3 19980529
US 1998-216070	A2 19981219
US 1999-414625	A1 19991008
US 2000-602980	B1 20000623
US 2000-664035	B1 20000918

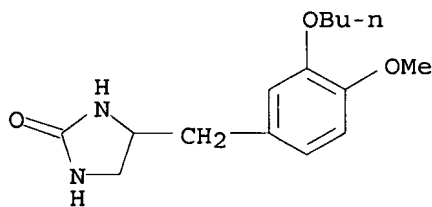
AB The invention provides pharmaceutical compns. containing compds. for the treatment of neoplasia in mammals. The phosphodiesterase inhibitory activity of a compound is determined along with cyclooxygenase inhibitory activity. Growth inhibitory and apoptosis inducing effects on cultured tumor cells are also determined. Compds. that exhibit phosphodiesterase inhibition, growth inhibition and apoptosis induction, but preferably not substantial prostaglandin inhibitory activity, are desirable for the treatment of neoplasia.

IT 29925-17-5, RO-20-1724

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)  
(antitumor agent identification methods, and pharmaceutical compns.)

RN 29925-17-5 CAPLUS

CN 2-Imidazolidinone, 4-[(3-butoxy-4-methoxyphenyl)methyl]- (CA INDEX NAME)



REFERENCE COUNT: 263 THERE ARE 263 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 6 OF 7 USPATFULL on STN

ACCESSION NUMBER: 1999:121379 USPATFULL

TITLE: Screening methods for cytokine inhibitors

INVENTOR(S): Mak, Vivian, Menlo Park, CA, United States

PATENT ASSIGNEE(S): Adolor Corporation, Malvern, PA, United States (U.S. corporation)

	NUMBER	KIND	DATE
PATENT INFORMATION:	US 5962477		19991005 <--
APPLICATION INFO.:	US 1998-97441		19980615 (9)
RELATED APPLN. INFO.:	Continuation-in-part of Ser. No. WO 1995-US4677, filed on 11 Apr 1995 which is a continuation-in-part of Ser. No. US 1995-400234, filed on 3 Mar 1995, now abandoned which is a continuation-in-part of Ser. No. US 1994-271287, filed on 6 Jul 1994, now abandoned which is a continuation-in-part of Ser. No. US 1994-225991, filed on 12 Apr 1994, now abandoned		
DOCUMENT TYPE:	Utility		
FILE SEGMENT:	Granted		
PRIMARY EXAMINER:	Tsang, Cecilia J.		
LEGAL REPRESENTATIVE:	Seidman, Stephanie L.Heller Ehrman White & McAuliffe		
NUMBER OF CLAIMS:	5		
EXEMPLARY CLAIM:	1		
NUMBER OF DRAWINGS:	6 Drawing Figure(s); 3 Drawing Page(s)		
LINE COUNT:	5138		

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention provides a number of screening methods for evaluating compounds capable of suppressing cytokine production either in vitro or in vivo. The methods generally involve stimulating the production of a cytokine in a cell, exposing a portion of the cells to a putative cytokine modulating agent and determining subsequent levels of cytokine production in the cells. Additionally, the present invention provides certain compounds identified by this method.

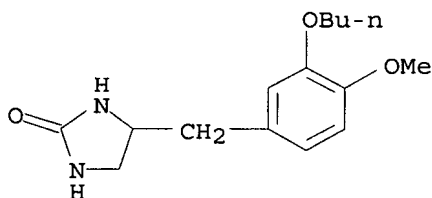
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 29925-17-5, RO 20-1724

(screening methods and formulations for cytokine inhibitors for treatment of inflammatory or immune conditions of skin)

RN 29925-17-5 USPATFULL

CN 2-Imidazolidinone, 4-[(3-butoxy-4-methoxyphenyl)methyl]- (CA INDEX NAME)



L6 ANSWER 7 OF 7 USPATFULL on STN

ACCESSION NUMBER: 1999:4360 USPATFULL  
TITLE: Method for identifying compounds for inhibition of cancerous lesions  
INVENTOR(S): Piazza, Gary A., Doylestown, PA, United States  
Pamukcu, Rifat, Spring House, PA, United States  
Thompson, W. Joseph, Mobile, AL, United States  
PATENT ASSIGNEE(S): Cell Pathways, Inc., Horsham, PA, United States (U.S. corporation)

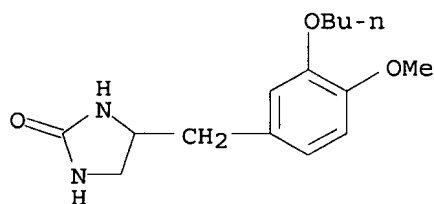
	NUMBER	KIND	DATE	
PATENT INFORMATION:	US 5858694		19990112	<--
APPLICATION INFO.:	US 1997-866027		19970530	(8)
DOCUMENT TYPE:	Utility			
FILE SEGMENT:	Granted			
PRIMARY EXAMINER:	Gitomer, Ralph			
LEGAL REPRESENTATIVE:	Brinks Hofer Gilson & Lione			
NUMBER OF CLAIMS:	26			
EXEMPLARY CLAIM:	1			
NUMBER OF DRAWINGS:	18 Drawing Figure(s); 13 Drawing Page(s)			
LINE COUNT:	1416			

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB This invention provides a method to identify compounds potentially useful for the treatment of neoplasia in mammals. The phosphodiesterase inhibitory activity of a compound is determined along with COX inhibitory activity. Growth inhibitory and apoptosis inducing effects on cultured tumor cells are also determined. Compounds that exhibit phosphodiesterase inhibition, growth inhibition and apoptosis induction, but not substantial prostaglandin inhibitory activity, are desirable for the treatment of neoplasia.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

IT 29925-17-5, Ro-20-1724  
(method evaluating inhibition of phosphodiesterase and cyclooxygenase activities, growth inhibition and apoptosis induction for identifying antineoplastic compds.)  
RN 29925-17-5 USPATFULL  
CN 2-Imidazolidinone, 4-[(3-butoxy-4-methoxyphenyl)methyl]- (CA INDEX NAME)



=> FIL STNGUIDE  
COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
64.21	72.67

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

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ENTRY	SESSION
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LAST RELOADED: Oct 19, 2007 (20071019/UP).

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(FILE 'HOME' ENTERED AT 11:19:52 ON 22 OCT 2007)

FILE 'REGISTRY' ENTERED AT 11:20:04 ON 22 OCT 2007

E "XX5"/CN 25

E "RO-1724"/CN 25

E "RO1724"/CN 25

E "RO-20-1724"/CN 25

E "4-(3-BUTOXY-4-METHOXYBENZYL)-2-IMIDAZOLIDINONE"/CN 25

L1 1 S E2

FILE 'MEDLINE, CAPLUS, WPIDS, USPATFULL' ENTERED AT 11:22:05 ON 22 OCT 2007

L2 875 S L1

L3 1 S L2 AND (CLL OR "CHRONIC LYMPHOCYTIC LEUKEMIA")

L4 140 S L1 AND (CANCER OR TUMOR)

L5 30 S L4 AND "LEUKEMIA"

L6 7 S L5 AND PY<2000

FILE 'STNGUIDE' ENTERED AT 11:33:50 ON 22 OCT 2007

=> s "PDE4" and "CLL"

0 "PDE4"

0 "CLL"

L7 0 "PDE4" AND "CLL"

=>

---Logging off of STN---

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Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS

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TOTAL  
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FULL ESTIMATED COST

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DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

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CA SUBSCRIBER PRICE

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